Yes

Yes

No

No





DECLARATION AND POWER OF ATTORNEY

(if only one name is listed below is claimed and for which a pate HUMAN PDGF BETA RECE Application Serial No. 08/258.) I have reviewed and understand referred to above. I acknowledg with Title 37, Code of Federal I	s and citizenship are as stated bel) or an original, first and joint in nt is sought on the invention en PTOR the specification of which 283 and was amended on the contents of the above identifie the duty to disclose information Regulations, Section 1.56. I claim s) for patent or inventor's certific	ventor (if plural inventors are notitled: INHIBITORY IMMU is attached hereto or (if applicable). ied specification, including the notice with the examplement of the examplemen	I am the original, first and sole inventor named below) of the subject matter which NOGLOBULIN POLYPEPTIDES TO X was filed on June 10, 1994 at a claims, as amended by any amendment in accordance for Title 35, United States Code, Section identified below any foreign application priority is claimed.
Prior Foreign Application(s)		••	
Country	Application No.	Date of Filing	Priority Claimed Under 35 USC 119

I claim the benefit under Title 35, United States Code, Section 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of Title 35, United States Code, Section 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, Section 1.56 which occurred between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Date of Filing	Status
Presently Unassigned	June 7, 1994	Patented X Pending Abandoned
07/801,795	December 2, 1991	Patented _X Pending Abandoned

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

William B. Smith, Reg. No. 30,223 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Reg. No. 36,429

Send Correspondence to: Karen B. Dow	Direct Telephone Calls to: (Name, Reg. No., telephone no.)
TOWNSEND and TOWNSEND KHOURIE and CREW	•
Steuart Street Tower	Karen B. Dow
One Market Plaza, 20th Floor	Reg. No. 29,684
San Francisco, CA 94105	(415) 326-2400, Ext. 217





			, 	
Full Name of Inventor 1	Last Name Ramakrishnan	First Name Vanitha	Middle Name or In	ițial
Residence & Citizenship	City Belmont	State/Foreign Country California	Country of Citizens India	hip
Post Office Address	Post Office Address 2019 Arbor Avenue	City Belmont	State/Country California	Zip Code 94002
Full Name of Inventor 2	Last Name Escobedo	First Name Maria	Middle Name or Initial Amelia	
Residence & Citizenship	City San Francisco	State/Foreign Country California	Country of Citizens Chile	hip
Post Office Address	Post Office Address 455 Upper Terrace #3	City San Francisco	State/Country California	Zip Code 94117
Full Name of Inventor 3	Last Name Fretto	First Name Larry	Middle Name or Initial J.	
Residence & . Citizenship	City Belmont	State/Foreign Country California	Country of Citizenship U.S.A.	
Post Office Address	Post Office Address 1553 Escondido Way	City Belmont	State/Country California	Zip Code 94002
Full Name of Inventor 4	Last Name Lokker	First Name Nathalie	Middle Name or Initial	
Residence & Citizenship	City San Francisco	State/Foreign Country California	Country of Citizenship The Netherlands	
Post Office Address	Post Office Address 741 40th Avenue	City San Francisco	State/Country California	Zip Code 94121
Full Name of Inventor 5	Last Name	First Name	Middle Name or Initial	
Residence & Citizenship	City	State/Foreign Country	Country of Citizenship	
Post Office Address	Post Office Address	City	State/Country	Zip Code
Full Name of Inventor 6	Last Name	First Name	Middle Name or Initial	
Residence & Citizenship	City	State/Foreign Country	Country of Citizenship	
Post Office Address	Post Office Address	City	State/Country	Zip Code

I further declare that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

Signature of Inventor 1 [autha/amaknshaan]	Signature of Inventor 2	Signature of Inventor 3
Vanitha Ramakrishnan	Maria Amelia Escobedo	Larry J. Fretto
Date 8 2 94	Date \$2/94	Date 8-2-94
Signature of Inventor 4 Nathalie Iokke	Signature of Inventor 5	Signature of Inventor 6
Date 7 8/2/94	Date	Date

K:\12418\18-2DEC.POA

(Page 3 of 3)

VER DE GATEMENT (DECLARATION) CLAIMING SWALL ENTITY STATUS (37 CFR 1.9(f) & 1.27(c)) - SMALL BUSINESS CONCERN

		an, Maria Amelia Escobedo, Larry J. Fretto and	Nathalie Lokker
Serial or Patent N Filed or Issued: J			
		POLYPEPTIDES TO HUMAN PDGF BETA RE	CEPTOR
I hereby declare t	that I am		•
	[] the owner of t	he small business concern identified below:	· ·
	= =	the small business concern empowered to act on	behalf of the concern identified below:
_			
	LL BUSINESS CONCERN		
ADDRESS OF S	SMALL BUSINESS CONCI	ERN 256 E. Grand Avenue, South San Francis	co, CA 94080
in 37 CFR 1.9(d) concern, including concern is the ave of the pay periods	, for purposes of paying red g those of its affiliates, does rage over the previous fisca of the fiscal year, and (2) of	duced fees to the United States Patent and Trader not exceed 500 persons. For purposes of this sta I year of the concern of the persons employed on	oncern as defined in 13 CFR 121.12, and reproduced nark Office, in that the number of employees of the tement, (1) the number of employees of the business a full-time, part-time or temporary basis during each irectly or indirectly, one concern controls or has the th.
to the invention, e		MUNOGLOBULIN POLYPEPTIDES TO HUMA	small business concern identified above with regard N PDGF BETA RECEPTOR by inventor(s) Vanitha
[]	the specification filed he	rewith	
[X]	application Serial No.	08/258,283 , filed June	10, 1994
[]	Patent No.	, issued	
1.9(d), or a nonpresent the status as small entry NAME	rofit organization under 37	CFR 1.9(e).	ot qualify as a small business concern under 37 CFR cation having rights to the invention averting to their
ADDRESS			
NAME	[] INDIVIDUAL	[] SMALL BUSINESS CONCERN	[] NONPROFIT ORGANIZATION
ADDRESS	[] INDIVIDUAL	[] SMALL BUSINESS CONCERN	[] NONPROFIT ORGANIZATION
prior to paying, or			resulting in loss of entitlement to small entity status ue after the date on which status as a small entity is
to be true; and fur or imprisonment, o	ther that these statements wor both, under section 1001	ere made with the knowledge that willful false sta	ements made on information and belief are believed tements and the like so made are punishable by fine h willful false statements may jeopardize the validity s directed.
NAME OF PERS	SON SIGNING	0.0.00 / 5.1.5	
	on if other than on	NER ROBER L SKIF	"Color of the Sale
	ERSON/SIGNING	256 E. Grand Avenue, South San Fran	ncisco, CA 94080
SIGNATURE	Ill	DATE	8/2/14
K:\12418\18-2.SES 39.MRG 5/95			



DECLARATION AND POWER OF ATTORNEY

As a below named inventor, I declare that:
My residence, post office address and citizenship are as stated below next to my name; I believe I am the original, first and sole inventor
(if only one name is listed below) or an original, first and joint inventor (if plural inventors are named below) of the subject matter which
is claimed and for which a patent is sought on the invention entitled: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO
HUMAN PDGF HETA RECEPTOR the specification of which is attached hereto or X was filed on June 10, 1994 at
Application Serial No. 08/258,283 and was amended on (if applicable).

I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above. I acknowledge the duty to disclose information which is material to the examination of this application in accordance with Title 37, Code of Federal Regulations, Section 1.56. I claim foreign priority benefits under Title 35, United States Code, Section 119 of any foreign applications(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of the application on which priority is claimed.

Prior Foreign Application(5)

Country	Application No.	Date of Filing	Priority Claimed Under 35 USC 119	
			Yes No	
			Yes No	

I claim the benefit under Title 35. United States Code, Section 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of Title 35, United States Code, Section 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, Section 1.56 which occurred between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Date of Filing	Status	
08/253,440	June 7, 1994	ParentedX Pending Abandoned	
07/801,795	December 2, 1991	Parented _X Pending Abandoned	

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

Shelley Precivale Eberle, Reg. No. 31,411 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Reg. No. 36,429

Send Correspondence to: Karen B. Dow	Direct Telephone Calls to: (Name, Reg. No., telephone co.)
TOWNSEND and TOWNSEND KHOURIE and CREW Steuart Street Tower One Market Plaza, 20th Floor San Francisco, CA 94105	Karen B. Dow Reg, No. 29,684 (415) 326-2400, Ext. 217

Full Name of Inventor 1	Last Name Ramakrishnan	First Name Vaniths	Middle Name or Initial	
Residence &	City Belmont	State/Foreign Country California	Country of Citizenship India	
Post Office Address	Post Office Address 2019 Arbor Avenue	City Belmont	State/Country California	Zip Code 94002
Full Name of Inventor 2	Last Name Escobedo	First Name Maria	Middle Name or In	nitial
Residence &	City San Francisco	State/Foreign Country California	Country of Citizenship Chile	
Post Office Address	Post Office Address 455 Upper Terrace #3	City San Francisco	State/Country California	Zip Code 94117
Full Name of Inventor 3	Last Name Fretto	First Name Larry	Middle Name or Initial J.	
Residence &	City Bekmont	State/Foreign Country California	Country of Citizenship U.S.A.	
Post Office Address	Post Office Address 1553 Escondido Way	City Belmont	State/Country California	Zip Code 94002

I further declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

Signature of Inventor 1	Signature of Inventor 2 Therewood. Esco Selfs	Signature of Inventor 3
Vanitha Ramakrishnan	Maria Amelia Escobedo	Larry J. Fretto
Date	Date 6/28/075	Date

(Page 2 of 2)

K-112419 IN-206C-2



DECLARATION AND POWER OF ATTORNEY

As a below named inventor, I declare that:
My residence, post office address and citizenship are as stated below next to my name; I believe I am the original, first and sole inventor
(if only one name is breed below) or an original, first and joint inventor (if plural inventors are named below) of the subject matter which
is claimed and for which a patent is sought on the invention entitled: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO
HUMAN PDGF BETA RECEPTOR the specification of which is attached hereto or _X was filed on June 10, 1994 as
Application Serial No. 08/258,283 and was amended on (if applicable).

I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above. I acknowledge the duty to disclose information which is material to the examination of this application in accordance with Title 37, Code of Federal Regulations, Section 1.56. I claim foreign priority benefits under Title 35, United States Code, Section 119 of any foreign applications(s) for patent or inventor's certificate listed below and have also identified below any foreign application for patent or inventor's certificate having a filing date before that of the application on which priority is claimed.

Prior Foreign Application(s)

Country	Application No.	Date of Filing	Priority Claimed Under 35 USC 119
			Yes No
			Yes No

I claim the benefit under Title 35. United States Code, Section 120 of any United States application(s) listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of Title 35, United States Code, Section 112, I acknowledge the duty to disclose material information as defined in Title 37, Code of Federal Regulations, Section 1.56 which occurred between the filing date of the prior application and the national or PCT international filing date of this application:

Application Serial No.	Date of Filing	Status
08/253,440	June 7, 1994	Patented _X Pending Abandoned
07/801.795	December 2, 1991	PatentedX Pending Abandoned

POWER OF ATTORNEY: As a named inventor, I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and transact all business in the Patent and Trademark Office connected therewith.

Shelley Precivale Eberle, Reg. No. 31,411 Karen B. Dow, Reg. No. 29,684 Randolph T. Apple, Reg. No. 36,429

Send Correspondence to:
Karen B. Dow
TOWNSEND and TOWNSEND KHOURIE and CREW
Steuart Street Tower
One Market Plaza, 20th Floor
San Francisco, CA 94105

Direct Telephone Calls to: (Name, Reg. No., telephone 20.)

Karen B. Dow Reg. No. 29,684 (415) 326-2400, Ext. 217

Full Name of Inventor 1	Last Name Ramakrishnan	First Name Vaniths	Middle Name or Initial	
Residence &	City Relmont	State/Foreign Country California	Country of Citizen India	ship
Post Office	Post Office Address 2019 Arbor Avenue	City Belmont	State/Country California	Zip Code 94002
Full Name of Inventor 2	Last Name Escobedo	First Name Maria	Middle Name or Initial Amelia	
Residence &	City San Francisco	State/Foreign Country California	Country of Citizenship Chile	
Post Office	Post Office Address 455 Upper Terrace #3	City San Francisco	State/Country California	Zip Code 94117
Full Name of Inventor 3	Last Name Fretto	First Name	Middle Name or Initial J.	
Residence & Citizenship	City Retmont	State/Foreign Country California	Country of Citizenship U.S.A.	
Post Office Address	Post Office Address 1553 Escondido Way	City Belmont	State/Country California	Zip Code 94002

I further declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

Signature of Inventor 1 () author Jamakur La	ar	Signature of Inventor 3
Vanitha Ramakrishnan	Maria Amelia Escobedo	Larry J. Fretto
	D.A.	Date 6.22-85
Date 22 JUNE 199512	Date	

(Page 2 of 2)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:)
Vanitha Ramakrishnan, et al.) Examiner: P. Gambel
Serial No.: 08/258,283) Art Unit: 1806
Filed: June 10, 1994)
For: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO HUMAN PDGF BETA RECEPTOR))) _)

Honorable Commissioner of Patents and Trademarks Washington, D.C. 20231

STATEMENT OF FACTS UNDER 37 C.F.R. 1.48(a)

Sir:

We, Vanitha Ramakrishnan, Maria Amelia Escobedo, Larry J. Fretto and Nathalie Lokker, hereby declare as follows:

- 1. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coinventors and co-applicants of patent application Serial No. 08/253, 440, filed on June 7, 1994 (the "Parent"), which is a file-wrapper-continuation of Serial No. 07/801,795, filed on December 2, 1991, describing immunoglobulin polypeptides that block binding of platelet- derived growth factor ("PDGF") to the platelet-derived growth factor receptor.
- 2. That the Parent describes an immunoglobulin polypeptide, referred to as the monoclonal antibody 2A1E2, and characterizes it as binding to the fifth domain of the extracellular domain of the PDGF receptor.
- 3. That the present application is a continuation-in-part of the Parent and characterizes 2A1E2 as binding to the second domain of the extracellular domain of the PDGF receptor, instead of the fifth domain as described in the Parent.

- 4. That Nathalie Lokker identified the second domain binding characteristics of 2A1E2. Based upon the fact that she provided additional data that was included in the present application and because claims were included to recite an immunoglobulin polypeptide the recognized an epitope in the second domain, Nathalie Lokker was named as an inventor at the time that the present application was filed.
- 5. It has now become apparent that Nathalie Lokker was incorrectly named as an inventor, through error and without any deceptive intention. Based upon our information and belief, the facts leading up to this discovery, are as follows:
- a) The issue of inventorship was raised by Examiner Gambel in the Office Action dated December 23, 1994. The Examiner took the position that since the Parent application described the same antibody that was being described in the present application, any differences in the epitope mapping (fifth vs. second Ig-like domain) only reflected a re-evaluation of the fine specificity of the same 2A1E2 antibody and did not change the inherent properties (i.e., structure and function) of the same deposited hybridoma (HB 10938). In light of this, the Examiner questioned why the inventive entities of the Parent and the present application were different.
- b) On June 1, 1995, in a telephonic interview between the Examiner and Applicants' representative Karen B. Dow, the Examiner more specifically questioned the inventorship status of Nathalie Lokker in view of the Examiner's assertion that the binding characteristics of the 2A1E2 antibody were inherent in the antibody, which was described in the Parent.
- c) On several occasions after this interview, Ms Dow discussed the inventorship issue with Applicants' attorneys, Carolyn R. Adler and Shelley P. Eberle, and it was determined that the binding characteristics of the monoclonal antibody 2A1E2 were inherent in the antibody, therefore that Nathalie Lokker was not an inventor of the subject matter as claimed in the present application.
- d) During the week of June 19, 1995, Ms. Eberle discussed this issue with us and we realized that an error in inventorship had been made and that it was appropriate to correct inventorship by the deletion of Nathalie Lokker as an inventor.
- 6. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coauthors of a publication entitled "A Novel Monoclonal Antibody Dependent on Domain 5 of the Platelet-Derived Growth Factor Beta Receptor Inhibits Ligand Binding and Receptor Activation", <u>Growth Factors</u>, (8):253-265 (1993); and that Joseph J. Seroogy, James E. Tomlinson and David L. Wolf are also named as co-authors on the publication. The contributions of the authors not named as inventors is as follows:
- a) David L. Wolf made the molecular biological construct and used it to transfect cells to produce the antigen (extracellular domain of the PDGF beta receptor), as described in column 1, lines 1-23, on page 255 of the publication. David L. Wolf also immunized rabbits to produce rabbit polyclonal serum, as described in column 1, lines 31-34 on page 255 of the publication.

- b) James E. Tomlinson characterized the antigen-expressing cell lines made by David L. Wolf, as described in column 1, lines 23-43 on page 255 of the publication. This antigen was later used as the immunogen for the production of the monoclonal antibodies. James E. Tomlinson also characterized the rabbit polyclonal antibodies made by David L. Wolf. These antibodies were used to determine the recombinant expression of the modified extracellular PDGF beta receptor by Western blot analysis, as described in column 1, lines 28-31 on page 255 of the publication.
- c) Joseph J. Seroogy performed the partial purification of the antigen (extracellular domain of the PDGF beta receptor) that was used to determine antibody titer in the ELISA, as described in column 2 on page 255 of the publication.
- d) It is our belief that co-authors Joseph J. Seroogy, James E. Tomlinson and David L. Wolf are not co-inventors of the claimed subject matter of the present application.

The undersigned declare further that all statements made herein of their own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements are made with the knowledge that willful false statements, and the like so made, are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issuing hereon.

Dated: <u>22 โบมะ 195</u>	Jani The Januakrishnan Vanitha Ramakrishnan
Dated:	Maria Amalia Fasahada
Dated: 6-22-85-	Maria Amelia Escobedo
Dated: <u>6-22-95</u>	Larry J. Fretto
	Nathalia Lokker

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:)
Vanitha Ramakrishnan, et al.) Examiner: P. Gambel
Serial No.: 08/258,283) Art Unit: 1806
Filed: June 10, 1994)
For: INHIBITORY IMMUNOGLOBULIN POLYPEPTIDES TO HUMAN PDGF BETA RECEPTOR))))

Honorable Commissioner of Patents and Trademarks Washington, D.C. 20231

STATEMENT OF FACTS UNDER 37 C.F.R. 1.48(a)

Sir.

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- 1. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coinventors and co-applicants of patent application Serial No. 08/253, 440, filed on June 7, 1994 (the "Parent"), which is a file-wrapper-continuation of Serial No. 07/801,795, filed on December 2, 1991, describing immunoglobulin polypeptides that block binding of platelet- derived growth factor ("PDGF") to the platelet-derived growth factor receptor.
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- 3. That the present application is a continuation-in-part of the Parent and characterizes 2A1E2 as binding to the second domain of the extracellular domain of the PDGF receptor, instead of the fifth domain as described in the Parent.

- 4. That Nathalie Lokker identified the second domain binding characteristics of 2A1E2. Based upon the fact that she provided additional data that was included in the present application and because claims were included to recite an immunoglobulin polypeptide the recognized an epitope in the second domain, Nathalie Lokker was named as an inventor at the time that the present application was filed.
- 5. It has now become apparent that Nathalie Lokker was incorrectly named as an inventor, through error and without any deceptive intention. Based upon our information and belief, the facts leading up to this discovery, are as follows:
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- b) On June 1, 1995, in a telephonic interview between the Examiner and Applicants' representative Karen B. Dow, the Examiner more specifically questioned the inventorship status of Nathalie Lokker in view of the Examiner's assertion that the binding characteristics of the 2A1E2 antibody were inherent in the antibody, which was described in the Parent.
- c) On several occasions after this interview, Ms Dow discussed the inventorship issue with Applicants' attorneys, Carolyn R. Adler and Shelley P. Eberle, and it was determined that the binding characteristics of the monoclonal antibody 2A1E2 were inherent in the antibody, therefore that Nathalie Lokker was not an inventor of the subject matter as claimed in the present application.
- d) During the week of June 19, 1995, Ms. Eberle discussed this issue with us and we realized that an error in inventorship had been made and that it was appropriate to correct inventorship by the deletion of Nathalie Lokker as an inventor.
- 6. That Vanitha Ramakrishnan, Maria Amelia Escobedo and Larry J. Fretto are the coauthors of a publication entitled "A Novel Monoclonal Antibody Dependent on Domain 5 of the Platelet-Derived Growth Factor Beta Receptor Inhibits Ligand Binding and Receptor Activation", Growth Factors, (8):253-265 (1993); and that Joseph J. Seroogy, James E. Tomlinson and David L. Wolf are also named as co-authors on the publication. The contributions of the authors not named as inventors is as follows:
- a) David L. Wolf made the molecular biological construct and used it to transfect cells to produce the antigen (extracellular domain of the PDGF beta receptor), as described in column 1, lines 1-23, on page 255 of the publication. David L. Wolf also immunized rabbits to produce rabbit polyclonal serum, as described in column 1, lines 31-34 on page 255 of the publication.



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The undersigned declare further that all statements made herein of their own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements are made with the knowledge that willful false statements, and the like so made, are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issuing hereon.

Dated:	
	Vanitha Ramakrishnan
Dated: 6/28/95	Maria Amelia Escobedo
	Maria Amelia Escobedo
Dated:	<u>.</u>
	Larry J. Fretto
Dated:	
•	Nathalie Lokker